

31/11 ATG TCC CAG AGC ACA CAG ACA AAT GAA TTC CTC AGT CCA GAG GTT TTC CAG CAT ATC TGG MSQSTQTNEFLSPEVFQHI 91/31 GAT TTT CTG GAA CAG CCT ATA TGT TCA GTT CAG CCC ATT GAC TTG AAC TTT GTG GAT GAA I D L N F V D E D F L E Q P I C S V Q P 151/51 CCA TCA GAA GAT GGT GCG ACA AAC AAG ATT GAG ATT AGC ATG GAC TGT ATC CGC ATG CAG EISMDCIRM P S E D G A T N K I 211/71 GAC TOG GAC CTG AGT GAC COC ATG TGG CCA CAG TAC ACG AAC CTG GGG CTC CTG AAC AGC рзргзррммьбллигеггй 271/91 ATG GAC CAG CAG ATT CAG AAC GGC TCC TCG TCC ACC AGT CCC TAT AAC ACA GAC CAC GCG STSPYNTDHA M_x D Q Q I Q N G S s 331/111 301/101 CAG AAC AGC GTC ACG GCG CCC TCG CCC TAC GCA CAG CCC AGC TCC ACC TTC GAT GCT CTC AQPSSTFDA QNSVTAPSP 391/131 TOT COA TOA COO GOO ATO COO TOO AAC ACO GAO TAC COA GGO COG CAC AGT TTO GAO GTG PSPAIPSNTDYPGPHSFD 451/151 421/141 TCC TTC CAG CAG TCG AGC ACC GCC AAG TCG GCC ACC TGG ACG TAT TCC ACT GAA CTG AAG YSTEL S F Q Q S S T A K S A T W T 511/171 ARA CTC TAC TGC CAR ATT GCA ARG ACA TGC CCC ATC CAG ATC ARG GTG ATG ACC CCA CCT 481/161 K L Y C Q I A K T C P I Q I K V M T P 571/191 CCT CAG GGA GCT GTT ATC CGC GCC ATG CCT GTC TAC AAA AAA GCT GAG CAC GTC ACG GAG P Q G A V I R A M P V Y K K A E H V T E 541/181 631/211 601/201 OTG OTG ANG COG TOC CCC ANC CAT GAG CTG AGC CGT GAA TTC AAC GAG GGA CAG ATT GCC V V K R C P N H E L S R E F N E G Q 691/231 CCT CCT AGT CAT TTG ATT CGA GTA GAG GGG AAC AGC CAT GCC CAG TAT GTA GAA GAT CCC 661/221 PPSHLIRVEGNSHAQY V E 751/251 ATC ACA GGA AGA CAG AGT GTG CTG GTA CCT TAT GAG CCA CCC CAG GTT GGC ACT GAA TTC PQVGT PYEP ITGRQSVLV 811/271 781/261 ACG ACA GTG TTG TAC AAT TTC ATG TGT AAC AGC AGT TGT GTT GGA GGG ATG AAC CGC Cgt TTVLYNFMCNSSCVGGMNRR 871/291 841/231 CCB ATT TTA ATC ATT GTT ACT CTG GAA ACC AGA GAT GGG CAA GTC CTG GGC CGA CGC TGC PILIIVTLETRDGQVLGRRC 931/311 901/301 TIT GAG GCC CGG ATC TOT GCT TGC CCA GGA AGA GAC AGG AAG GCG GAT GAA GAT AGC ATC FEARICACPGRDRKADED 991/331 961/321 AGA AAG CAG CAA GTT TOG GAC AGT ACA AAG AAC GGT GAT GGT ACG AAG CGC CCG TTT CGT Q Q V S D S T K N G D G T K R R K 1051/351 1021/341 CAG AAC ACA CAT GGT ATC CAG ATG ACA TCC ATC AAG AAA CGA AGA TCC CCA GAT GAT GAA H G I Q M T S I K K R R S P D D 1111/371 CTG TTA TAC TTA CCA GTG AGG GGC CGT GAG ACT TAT GAA ATG CTG TTG AAG ATC AAA GAG LLYLPVRGRETYEMLLKIKE 1171/391 TCC CTG GAA CTC ATG CAG TAC CTT CCT CAG CAC ACA ATT GAA ACG TAC AGG CAA CAG CAA SLELMQYLPQHTIETYRQQQQ

1231/411 CAG CAG CAG CAC CAG CAC TTA CAG AAA CAG ACC TCA ATA CAG TCT CCA TCA TAT QQHQHLDQKQTSIQ S Y 1291/431 GOT AAC AGC TCC CCA CCT CTG AAC AAA ATG AAC AGC ATG AAC AAG CTG CCT TCT GTG AGC с и s s p р г и к и и s и и к г р 1351/451 CAG CTT ATC AAC CCT CAG CAG CGC AAC GCC CTC ACT CCT ACA ACC ATT CCT GAT GGC ATG QLINPQQRNALTPTIPD GGA GCC AAC ATT CCC ATG ATG GGC ACC CAC ATG CCA ATG GCT GGA GAC ATG AAT GGA CTC G A N I P M M G T H M P M A G D AGO COO ACO CAG GOA CTC CCT CCC CCA CTC TCC ATG CCA TCC ACC TCC CAC TGC ACA CCC S M P S T SPTQALPPPL 1531/511 CCA CCT CCG TAT CCC ACA GAT TGC AGC ATT GTC AGT TTC TTA GCG AGG TTG GGC TGT TCA v s F PPPYPTDCSI 1591/531 TCA TOT CTG GAC TAT TTC ACG ACC CAG GGG CTG ACC ACC ATC TAT CAG ATT GAG CAT TAC S C L D Y F T T Q G L T T TCC ATG GAT GAT CTG GCA AGT CTG AAA ATC CCT GAG CAA TTT CGA CAT GCG ATC TGG AAG S M D D L A S L K I P E Q F R H A I W 1711/571 GGC ATC CTG GAC CAC CGG CAG CTC CAC GAA TTC TCC TCC CCT TCT CAT CTC CTG CGG ACC G I L D H R Q L H E F S S 1771/591 CCA AGC AGT GCC TCT ACA GTC AGT GTG GGC TCC AGT GAG ACC CGG GGT GAG CGT GTT ATT PSSASTVSVGSSETRGERV 1831/611 GAT GCT GTG CGA TTC ACC CTC CGC CAG ACC ATC TCT TTC CCA CCC CGA GAT GAG TGG AAT DAVRFTLRQTISFPPRDE 1891/631 GAC TTC AAC TTT GAC ATG GAT GCT CGC CGC AAT AAG CAA CAG CGC ATC AAA GAG GAG GGG F N F D M D A R R N K Q Q R I K E 1921/641 GAG TGA

31/11 ATG TCC CAG AGC ACA CAG ACA AAT GAA TTC CTC AGT CCA GAG GTT TTC CAG CAT ATC TGG M S Q S T Q T N E F L S P E V F Q H I 91/31 GAT TIT CTG GAA CAG CCT ATA TGT TCA GTT CAG CCC ATT GAC TIG AAC TTT GTG GAT GAA QPIDLN DFLEQPICS V 151/51 CCA TCA GAA GAT GGT GCG ACA AAC AAG ATT GAG ATT AGC ATG GAC TGT ATC CGC ATG CAG EISMDCIR PSEDGATNKI 211/71 CAC TOG GAC CTG AGT GAC CCC ATG TGG CCA CAG TAC ACG AAC CTG GGG CTC CTG AAC AGC QYTNLGL SDLSDPMWP 271/91 ATG GAC CAG CAG ATT CAG AAC GGC TCC TCG TCC ACC AGT CCC TAT AAC ACA GAC CAC GCG S T S MDQQIQNGSS Y N T 331/111 CAG AAC AGC GTC ACG GCG CCC TCG CCC TAC GCA CAG CCC AGC TCC ACC TTC GAT GCT CTC 301/101 S S T Q N S V T A P S P Y A Q P 391/131 361/121 TOT COA TOA COO GOO ATO COO TOO AAC ACO GAO TAD COA GGO COG CAC AGT TTO GAO GTG PSPAIPSNTDYPGPH s F 451/151 421/141 TCC TTC CAG CAG TCG AGC ACC GCC AAG TCG GCC ACC TGG ACG TAT TCC ACT GAA CTG AAG YSTELK S F Q Q S S T A K S A T W 511/171 481/161 AAA CTC TAC TGC CAA ATT GCA AAG ACA TGC CCC ATC CAG ATC AAG GTG ATG ACC CCA CCT K L Y C Q I A K T C P I Q I K V M T 571/191 541/181 CCT CAG GGA GCT GTT ATC CGC GCC ATG CCT GTC TAC AAA AAA GCT GAG CAC GTC ACG GAG V Y K K A E H V T PQGAVIRAMP 631/211 GTG GTG AAG CGG TGC CCC AAC CAT GAG CTG AGC CGT GAA TTC AAC GAG GGA CAG ATT GCC V V K R C P N H E L S R E FNEGQ 691/231 661/221 CCT CCT AGT CAT TTG ATT CGA GTA GAG GGG AAC AGC CAT GCC CAG TAT GTA GAA GAT CCC P P S H L I R V E G N S H A Q Y V E D 751/251 721/241 ATC ACA GGA AGA CAG AGT GTG CTG GTA CCT TAT GAG CCA CCC CAG GTT GGC ACT GAA TTC YEPPQ ITGRQSVLVP 811/271 781/261 ACG ACA GTC TTG TAC AAT TTC ATG TGT AAC AGC AGT TGT GTT GGA GGG ATG AAC CGC CGL TTVLYNFMCNSSCVGGMN 871/291 CCa att tha atc att git act ctg gaa acc aga gat ggg caa gtc ctg ggc cga cgc tgc PILIVTLE T R D G Q V L G R R C 931/311 901/301 TTT GAG GCC CGG ATC TGT GCT TGC CCA GGA AGA GAC AGG AAG GCG GAT GAA GAT AGC ATC FEARICACPGRDRKADED 991/331 961/321 AGA AAG CAG CAA GTT TCG GAC AGT ACA AAG AAC GGT GAT GGT ACG AAG CGC CCG TTT CGT GTKRPFR RKQQVSDSTKNGD 1051/351 1021/341 CAG AAC ACA CAT GGT ATC CAG ATG ACA TCC ATC AAG AAA CGA AGA TCC CCA GAT GAT GAA NTHGIQMTSIKKRRSPD 1111/371 1081/361 CTG TTA TAC TTA CCA GTG AGG GGC CGT GAG ACT TAT GAA ATG CTG TTG AAG ATC AAA GAG TYE'ML L LLYLPVRGRE 1171/391 1141/381 TCC CTG GAA CTC ATG CAG TAC CTT CCT CAG CAC ACA ATT GAA ACG TAÇ AGG CAA CAG CAA S L E L M Q Y L P Q H T I E T Y R Q Q Q

1231/411 1201/401 CAG CAG CAG CAC CAG CAC TTA CAG AAA CAG ACC TCA ATA CAG TCT CC CT TCA TAT Q Q Q H Q H L L Q K Q T S I Q S P S S Y 1291/431 1261/421 GGT AAC AGC TCC CCA CCT CTG AAC AAA ATG AAC AGC ATG AAC AAG CTG CCT TCT GTG AGC G N S S P P L N K M N S M N K L P S V S 1351/451 1321/441 CAG CTT ATC AAC CCT CAG CAG CGC AAC GCC CTC ACT CCT ACA ACC ATT CCT GAT GGC ATG Q L I N P Q Q R N A. L T P T T I P D G M 1411/471 1381/461 GGA GCC AAC ATT CCC ATG ATG GGC ACC CAC ATG CCA ATG GCT GGA GAC ATG AAT GGA CTC
G A N I P M M G T H M P M A G D M N G L 1441/481 1471/491 AGC CCC ACC CAG GCA CTC CCT CCC CCA CTC TCC ATG CCA TCC ACC TCC CAC TGC ACA CCC SPTQALPPPLSMPSTSHCTP-1501/501 1531/511 CCA CCT CCG TAT CCC ACA GAT TGC AGC ATT GTC AGG ATC TGG CAA GTC TGA PPPYPTDCSIVRIWQV*

31/11 ATG TCC CAG AGC ACA CAG ACA AAT GAA TTC CTC AGT CCA GAG GTT TTC CAG CAT ATC TGG M S Q S T Q T N E F L S P E V F Q H I 91/31 61/21 GAT TIT CTG GAA CAG CCT ATA TGT TCA GTT CAG CCC ATT GAC TTG AAC TTT GTG GAT GAA DLNFVDE D F L E Q P I C S V Q P I 151/51 121/41 CCA TCA GAA GAT GGT GCG ACA AAC AAG ATT GAG ATT AGC ATG GAC TGT ATC CGC ATG CAG PSEDGATNKIEISMDCIRM 211/71 GAC TCG GAC CTG AGT GAC CCC ATG TGG CCA CAG TAC ACG AAC CTG GGG CTC CTG AAC AGC QYTNLGLLNS D S D L S D P M W P 271/91 241/81 ATG GAC CAG CAG ATT CAG AAC GGC TCC TCG TCC ACC AGT CCC TAT AAC ACA GAC CAC GCG M D Q Q I Q N G S S T S P Y N T D H 331/111 301/101 CAG AAC AGC GTC ACG GCG CCC TCG CCC TAC GCA CAG CCC AGC TCC ACC TTC GAT GCT CTC Q N S V T A P S P Y A Q P S S T F D A L 391/131 361/121 TCT CCA TCA CCC GCC ATC CCC TCC AAC ACC GAC TAC CCA GGC CCG CAC AGT TTC GAC GTG SPSPAIPSNT D Y P G P H S 451/151 421/141 TCC TTC CAG CAG TCG AGC ACC GCC AAG TCG GCC ACC TGG ACG TAT TCC ACT GAA CTG AAG S F Q Q S S T A K S A T W T Y S T E L K 511/171 ANA CTC TAC TGC CAA ATT GCA AAG ACA TGC CCC ATC CAG ATC AAG GTG ATG ACC CCA CCT K L Y C Q I A K T C P I Q I K V M T P P 571/191 541/181 CCT CAG GGA GCT GTT ATC CGC GCC ATG CCT GTC TAC AAA AAA GCT GAG CAC GTC ACG GAG E H V PQGAVIRAMPVYKKA 601/201 631/211 GTG GTG AAG CGG TGC CCC AAC CAT GAG CTG AGC CGT GAA TTC AAC GAG GGA CAG ATT GCC V V K R C P N H E L S R E F N E G Q 661/221 691/231 CCT CCT AGT CAT TTG ATT CGA GTA GAG GGG AAC AGC CAT GCC CAG TAT GTA GAA GAT CCC P P S H L I R V E G N S H A Q Y V E D 721/241 751/251 ATC ACA GGA AGA CAG AGT GTG CTG GTA CCT TAT GAG CCA CCC CAG GTT GGC ACT GAA TTC TGRQSVLVP Y E P V G P 0 811/271 ACG ACA GTC TTG TAC AAT TTC ATG TGT AAC AGC AGT TGT GGT GGA GGG ATG AAC CGC CGL T T V L Y N F M C N S S C V G G M N R R 871/291 CCA aTT TTA ATC ATT GTT ACT CTG GAA ACC AGA GAT GGG CAA GTC CTG GGC CGA CGC TGC PILIIVTLETRDG QVLGRRC 901/301 931/311 TTT GAG GCC CGG ATC TGT GCT TGC CCA GGA AGA GAC AGG AAG GCG GAT GAA GAT AGC ATC F E A R I C A C P G R D R K A D E D S 961/321 991/331 AGA AAG CAG CAA OTT TCG GAC AGT ACA AAG AAC GGT GAT GGT ACG AAG CGC CCG TTT CGT RKQQVSDSTKNGDGTKRPF 1021/341 1051/351 CAG AAC ACA CAT GGT ATC CAG ATG ACA TCC ATC AAG AAA CGA AGA TCC CCA GAT GAT GAA NTHGIQMTS IKKRRSPDD 1081/361 1111/371 CTG TTA TAC TTA CCA GTG AGG GGC CGT GAG ACT TAT GAA ATG CTG TTG AAG ATC AAA GAG L P V R G R E T Y E M L L K I K E 1171/391 1141/381 TCC CTG GAA CTC ATG CAG TAC CTT CCT CAG CAC ACA ATT GAA ACG TAC\AGG CAA CAG CAA г г г и б л г в б н т г в т л

1201/401

CAG CAG CAG CAC CAG CAC TT T CAG AAA CAT CTC CTT TCA GCC TGC T GAG AAT GAG Q Q Q H Q H L L Q K H L L S A C F R N E 1291/431

CTT GTG GAG CCC CGG AGA GAA ACT CCA AAA CAA TCT GAC GTC TTC TTT AGA CAT TCC AAG L V E P R R E T P K Q S D V F F R H S K 1321/441

CCC CCA AAC CGA TCA GTG TAC CCA TAG P P N R S V Y P *

1/1 31/11 ATG TTG TAC CTG GAA AAC AAT GCC CAG ACT CAA TTT AGT GAG CCA CAG TAC ACG AAC CTG MLYLENNAQT Q F SEPQYTNL 61/21 91/31 GGG CTC CTG AAC AGC ATG GAC CAG CAG ATT CAG AAC GGC TCC TCG TCC ACC AGT CCC TAT G L L N S M D Q Q I Q N G S S T S P Ī51/S1 AAC ACA GAC CAC GCG CAG AAC AGC GTC ACG GCG CCC TCG CCC TAC GCA CAG CCC AGC TCC N T D H A Q N S V T A P S P Y A Q P S S 211/71 ACC TTC GAT GCT CTC TCT CCA TCA CCC GCC ATC CCC TCC AAC ACC GAC TAC CCA GGC CCG T F D A L S P S P A I P S N T D Y P G 241/81 271/91 CAC AGT TTC GAC GTG TCC TTC CAG CAG TCG AGC ACC GCC AAG TCG GCC ACC TCG ACG TAT H S F D V S F Q Q S S T A K S A T 301/101 331/111 TCC ACT GAA CTG AAG AAA CTC TAC TGC CAA ATT GCA AAG ACA TGC CCC ATC CAG ATC AAG S T E L K K L Y C Q I A K T C P I Q I K 361/121 391/131 GTG ATG ACC CCA CCT CCT CAG GGA GCT GTT ATC CGC GCC ATG CCT GTC TAC AAA AAA GCT VMTPPPQGAV I R A M P 421/141 451/151 GAG CAC GTC ACG GAG GTG GTG AAG CGG TGC CCC AAC CAT GAG CTG AGC CGT GAA TTC AAC 481/161 511/171 AGG GGA CAG ATT GCC CCT CCT AGT CAT TTG ATT CGA GTA GAG GGG AAC AGC CAT GCC CAG E G Q I A P P S H L I R V EGNSHAQ 541/181 571/191 TAT OTA GAA GAT CCC ATC ACA GGA AGA CAG AGT GTG CTG GTA CCT TAT GAG CCA CCC CAG HAY V E D P I T G R Q S V L V P 601/201 631/211 GTT GGC ACT GAA TTC ACG ACA GTC TTG TAC AAT TTC ATG TGT AAC AGC AGT TGT GTA GGA V G T E F T T V L Y N F M C N S S C V 691/231 GGG ATG AAC CGC CGE CCA ATT TTA ATC ATT GTT ACT CTG GAA ACC AGA GAT GGG CAA GTC G M N R R P I L I I V T L E T R D G Q V 721/241 751/251 THE CTG GGC CGA CGC TGC TTT GAG GCC CGG ATC TGT GCT TGC CCA GGA AGA GAC AGG AAG GCG L G R R C F E A R I C A C P G R D R 781/261 GAT GAA GAT AGC ATC AGA AAG CAG CAA GTT TCG GAC AGT ACA AAG AAC GGT GAT GGT ACG 811/271 D E D S I R K Q Q V S D S T K N G D 871/291 AAG CGC CCG TTT CGT CAG AAC ACA CAT GGT ATC CAG ATG ACA TCC ATC AAG AAA CGA AGA K R P F R Q N T H G I Q M T s IKKRR 901/301 931/311 TCC CCA GAT GAT GAA CTG TTA TAC TTA CCA GTG AGG GGC CGT GAG ACT TAT GAA ATG CTG D D E L L Y L P V R G RETYEM 961/321 991/331 TTG AAG ATC AAA GAG TCC CTG GAA CTC ATG CAG TAC CTT CCT CAG CAC ACA ATT GAA ACG K I K E S L E L M Q Y L P Q H T 1021/341 1051/351 TAC AGG CAA CAG CAG CAG CAG CAC CAG CAC TTA CTT CAG AAA CAG ACC TCA ATA CAG R Q Q Q Q Q H Q H L L Q K Q T S I Q 1081/361 1111/371 TOT COA TOT TOA TAT GOT AAC AGO TOO COA COT CTG AAC AAA ATG AAC AGO ATG AAC AAG S P S S Y G N S S P P L N K M N S M N 1141/381 1171/391 CTG CCT TCT GTG AGC CAG CTT ATC AAC CCT CAG CAG CGC AAC GCC CTC ACT CCT ACA ACC L P S V S Q L I N P Q Q R N A L T P

1231/411 1201/401 ATT CCT GAT GGC ATG GGA GCC ... ATT CCC ATG ATG GGC ACC CAC ATG CCA ... G GCT GGA I P D G M G A N I P M M G T H M P M A G 1291/431 1261/421 GAC ATG AAT GGA CTC AGC CCC ACC CAG GCA CTC CCT CCC CCA CTC TCC ATG CCA TCC ACC D M N G L S P T Q A L P P P L S M P 1351/451 1321/441 TCC CAC TGC ACA CCC CCA CCT CCG TAT CCC ACA GAT TGC AGC ATT GTC AGT TTC TTA GCG S H C T P P P P Y P T D C S I V S F L A 1381/461 1411/471 AGG TTG GGC TGT TCA TCA TGT CTG GAC TAT TTC ACG ACC CAG GGG CTG ACC ACC ATC TAT FTTQGL T T RLGCSSCLDY 1471/491 1441/481 CAG ATT GAG CAT TAC TOC ATG GAT GAT CTG GCA AGT CTG AAA ATC CCT GAG CAA TTT CGA Q I E H Y S M D D L A S L K I P E Q F R 1501/501 1531/511 CAT GCG ATC TGG AAG GGC ATC CTG GAC CAC CGG CAG CTC CAC GAA TTC TCC TCC CCT TCT HAIWKGILDHRQLHEFSSPS 1591/531 1561/521 CAT CTC CTG CGG ACC CCA AGC AGT GCC TCT ACA GTC AGT GTG GGC TCC AGT GAG ACC CGG H L L R T P S S A S T V S V G S 1651/551 1621/541 GGT GAG CGT GTT ATT GAT GCT GTG CGA TTC ACC CTC CGC CAG ACC ATC TCT TTC CCA CCC G E R V I D A V R F T L R Q T I S F P P 1681/561 1711/571 CGA GAT GAG TGG AAT GAC TTC AAC TTT GAC ATG GAT GCT CGC CGC AAT AAG CAA CAG CGC RDEWNDFNFDMDARNKO 1741/581 ATC AAA GAG GAG GAG TGA IKEEGE

31/11 1/1 ATG TTG TAC CTG GAA AAC AAT GCC CAG ACT CAA TTT AGT GAG CCA CAG TAC ACG AAC CTG Y L E N N A Q T Q F S E P Q Y T 91/31 61/21 GGG CTC CTG AAC AGC ATG GAC CAG CAG ATT CAG AAC GGC TOC TOG TOC ACC AGT CCC TAT SS G L L N S M D Q Q I Q N G S T s P 151/51 AAC ACA GAC CAC GCG CAG AAC AGC GTC ACG GCG CCC TOG CCC TAC GCA CAG CCC AGC TCC N T D H A Q N S V T A P S P Y A Q P S 211/71 181/61 ACC TTC GAT GCT CTC TCT CCA TCA CCC GCC ATC CCC TCC AAC ACC GAC TAC CCA GGC CCG T F D A L S P S P A I P S N T D Y 271/91 CAC AGT TTC GAC GTG TCC TTC CAG CAG TOG AGC ACC GCC AAG TCG GCC ACC TGG ACG TAT H S F D V S F Q Q S S T A K S A T W T Y 331/111 301/101 TCC ACT GAA CTG AAG AAA CTC TAC TGC CAA ATT GCA AAG ACA TGC CCC ATC CAG ATC AAG STELKKLYCQIAKTCPIQ 391/131 361/121 GTG ATG ACC CCA CCT CCT CAG GGA GCT GTT ATC CGC GCC ATG CCT GTC TAC AAA AAA GCT V M T P P P Q G A V I R A M P V Y K K A 421/141 451/151 GAG CAC GTC ACG GAG GTG GTG AAG CGG TGC CCC AAC CAT GAG CTG AGC CGT GAA TTC AAC EHVTEVVKRCPNHELSRE 511/171 481/161 GAG GGA CAG ATT GCC CCT CCT AGT CAT TTG ATT CGA GTA GAG GGG AAC AGC CAT GCC CAG IRVEGNSHAQ EGQIAPPSHL 571/191 TAT GTA GAA GAT CCC ATC ACA GGA AGA CAG AGT GTG CTG GTA CCT TAT GAG CCA CCC CAG YVEDPITGRQSVLVPYEP P O 601/201 631/211 GTT GGC ACT GAA TTC ACG ACA GTC TTG TAC AAT TTC ATG TOT AAC AGC AGT TGT GTA GGA V G T E F T T V L Y N F M C N S S C V G 661/221 691/231 GGG ATG AAC CGC CGt CCa aTT TTA ATC ATT GTT ACT CTG GAA ACC AGA GAT GGG CAA GTC D G G M N R R P I L I I V T L E T R 721/241 751/251 CTG GGC CGA CGC TGC TTT GAG GCC CGG ATC TGT GCT TGC CCA GGA AGA GAC AGG AAG GCG LGRRCFEARICACPGRDRKA 781/261 811/271 GAT GAA GAT AGC ATC AGA AAG CAG CAA GTT TCG GAC AGT ACA AAG AAC GGT GAT GGT ACG SIRKQQVSDSTKNGDGT 871/291 AAG OGC COG TIT COT CAG AAC ACA CAT GOT ATC CAG ATG ACA TCC ATC AAG AAA CGA AGA KRPFRQNTHG IQMT SIKKRR 931/311 TCC CCA GAT GAT GAA CTG TTA TAC TTA CCA GTG AGG GGC COT GAG ACT TAT GAA ATG CTG SPDDELLYLPVRGRETYEML 961/321 991/331 TTG AAG ATC AAA GAG TCC CTG GAA CTC ATG CAG TAC CTT CCT CAG CAC ACA ATT GAA ACG LKIKESLELMQYLP QHTIET 1021/341 1051/351 TAC AGG CAA CAG CAA CAG CAG CAC CAG CAC TIA CTT CAG AAA CAG ACC TCA ATA CAG Y R Q Q Q Q Q H Q H L L Q K Q T S I Q 1081/361 1111/371 TCT CCA TCT TCA TAT GGT AAC AGC TCC CCA CCT CTG AAC AAA ATG AAC AGC ATG AAC AAG SYGNSSPPLNKMNSMN 1141/381 1171/391 CTG OCT TCT GTG AGC CAG CTT ATC AAC CCT CAG CAG CGC AAC GCC CTC ACT CCT ACA ACC L P S V S Q L I N P Q Q R N A L T P T T

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1/1
                            31/11
MLYLENNAQTQF
                                  SEPQYTNL
61/21
                            91/31
GGG CTC CTG AAC AGC ATG GAC CAG CAG ATT CAG AAC GGC TCC TCG TCC ACC AGT CCC TAT
G L L N S M D Q Q I
                            Q N
                                 G S
                                       S S
                            151/51
AAC ACA GAC CAC GCG CAG AAC AGC GTC ACG GCG CCC TCG CCC TAC GCA CAG CCC AGC TCC
NTDHAQNSVT
                            APSPYAQP
                            211/71
ACC TTC GAT GCT CTC TCT CCA TCA CCC GCC ATC CCC TCC AAC ACC GAC TAC CCA GGC CCG
TFDALSPSPAIPSNTDYPGP
241/31
                            271/91
CAC AGT TTC GAC GTG TCC TTC CAG CAG TCG AGC ACC GCC AAG TCG GCC ACC TGG ACG TAT
H S F D V S F Q Q S
                            301/101
                            331/111
TCC ACT GAA CTG AAG AAA CTC TAC TGC CAA ATT GCA AAG ACA TGC CCC ATC CAG ATC AAG
STELKKLYCQ
                            IAKTCPIQIK
361/121
                            391/131
OTG ATG ACC CCA CCT CCT CAG GGA GCT GTT ATC CGC GCC ATG CCT GTC TAC AAA AAA GCT
VMTPPPQGAV
                           IRAMPVYKKA
421/141
                            451/151
GAG CAC GTC ACG GAG GTG GTG AAG CGG TGC CCC AAC CAT GAG CTG AGC CGT GAA TTC AAC
E H V T E V V K R C
                           PNHELSREF
481/161
                            511/171
GAG GGA CAG ATT GCC CCT CCT AGT CAT TTG ATT CGA GTA GAG GGG AAC AGC CAT GCC CAG
EGQIAPPSHL
                            I R V
                                    EGNSHAQ
541/191
                            571/191
TAT GTA GAA GAT CCC ATC ACA GGA AGA CAG AGT GTG CTG GTA CCT TAT GAG CCA CCC CAG
YVEDPITGRQ
                           SVLVPYEPPQ
601/201
                            631/211
GIT GGC ACT GAA TIC ACG ACA GIC TIG TAC AAT TIC AIG TGI AAC AGC AGI TGI GIA GGA
V G T E F T T V L Y N F M C N S S C
661/221
                            691/231
GGG ATG AAC CGC CGt CCA att tta atc att git act ctg gaa acc aga gat ggg caa gtc
GMNRRPILIIVTL
                                   ETRDG
                            751/251
CTG GGC CGA CGC TGC TTT GAG GCC CGG ATC TGT GCT TGC CCA GGA AGA GAC AGG AAG GCG
     RRCFEARICACPGRDRKA
781/261
                            811/271
gat gaa gat agc atc aga aag cag caa git tog gac agt aca aag aac ggt gat ggt acg
D E D S I R K Q Q V S D S
                                   TKNGDGT
841/281
                            871/291
AAG CGC CCG TIT CGT CAG AAC ACA CAT GGT ATC CAG ATG ACA TCC ATC AAG AAA CGA AGA
K R P F R Q N T H G
                            IQMTSIKKR
901/301
                            931/311
TCC CCA GAT GAT GAA CTG TTA TAC TTA CCA GTG AGG GGC CGT GAG ACT TAT GAA ATG CTG
SPDDELLYP
                           V R G R E T Y E M L
961/321
                            991/331
TTG ANG ATC ANA GAG TCC CTG GAA CTC ATG CAG TAC CTT CCT CAG CAC ACA ATT GAA ACG
       K E S L E L M Q Y L
                                      QHTIET
1021/341
                            1051/351
TAC AGG CAA CAG CAG CAG CAG CAG CAC TTA CTT CAG AAA CAT CTC CTT TCA GCC
       Q Q Q Q H Q H L L Q K H L L S
                            1111/371
TGC TTC AGG AAT GAG CTT GTG GAG CCC CGG AGA GAA ACT CCA AAA CAA TCT GAC GTC TTC
       NELVEPRRETYPKQSDVF
1141/381
                            1171/391
TTT AGA CAT TCC AAG CCC CCA AAC CGA TCA GTG TAC CCA TAG
F R H S K P P N R S V Y P
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1/1 31/11 ATG AAT TIT GAA ACT TOA CGG TOT GCC ACC CTA CAG TAC TGC CCC GAC CCT TAC ATC CAG MNFETSRCATLQ YCPDPYI 91/31 CGT TTC ATA GAA ACC CCA GCT CAT TTC TCG TGG AAA GAA AGT TAT TAC AGA TCT GCC ATG I E T PAHFS W K E S Y Y R S 151/51 TOG CAG AGC ACC CAG ACA AGC GAG TTC CTC AGC CCA GAG GTC TTC CAG CAT ATC TGG GAT S Q S T Q T. S E F L SPEVFQHIW 181/61 211/71 TIT CIG GAA CAG CCT ATA TGC TCA GIA CAG CCC ATC GAG TIG AAC TIT GIG GAT GAA CCT FL EQFICSVQPIELNFVDE 241/81 271/91 TCC GAA AAT GGT GCa aCa aaC aaG ATT GAG ATT AGC ATG GAT TGT ATC CGC ATG Caa GAC S E NGATNKIE I S M D C I R M 301/101 331/111 TCA GAG CTC AGT GAG CCC ATG TGG CCA CAG TAC ACG AAC CTG GGG CTC CTG AAC AGC ATG S. D. L. S. D. P. M. W. P. Q. Y T N L G L L N S M 361/121 391/131 GAC CAG CAG ATT CAG AAC GGC TCC TCG TCC ACC AGC CCC TAC AAC ACA GAC CAC GCA CAG Q Q I Q N G S S S T S P Y N T D H A Q 421/141 451/151 AAT AGC GTG AGG GCG CCC TGG CCC TAT GCA CAG CCC AGC TCC ACC TTT GAT GCC CTC TCT N S V T A P S P Y A Q P S S T F D A 481/161 511/171 CCA TCC CCT GCC ATT CCC TCC AAC ACA GAT TAC CCG GGC CCA CAC AGC TTC GAT GTG TCC PSPAIPSNTDYPG 541/191 571/191 TTC CAG CAG TCA AGC ACT GCC AAG TCA GCC ACC TGG ACG TAT TCC ACC GAA CTG AAG AAG FQQSSTAKSATWT YSTELKK 601/201 631/211 CTG TAC TGC CAG ATT GCG AAG ACA TGC CCC ATC CAG ATC AAG GTG ATG ACC CCA CCC CCA YCQIAKTCPIQI KVMTP 661/221 691/231 CAG GGC GCT GTE ATC CGT GCC ATG CCT GTC TAC AAG AAA GCT GAG CAT GTC ACC GAG GTT GAVIRAMPV YKKAEHVTEV 751/251 STG AAA CGA TGC CCT AAC CAT GAG CTG AGC CGT GAG TTC AAT GAG GGA CAG ATT GCC CCT V K R C P N H E L S R E F N E G Q I A 811/271 CCC AGT CAT CTG ATT CGA GTA GAA GGG AAC AGC CAT GCC CAG TAT GTA GAA GAT CCT ATC SHLIRVEGNSHAQY(VEDPI 841/281 871/291 ACG GGA AGG CAG AGC GTG CTG GTC CCT TAT GAG CCA CCA CAG GTT GGC ACT GAA TTC ACA G R Q S V L V P Y E P P Q V G 901/301 931/311 ACA GTC CTG TAC BAT TTC ATG TGT AAC AGC AGC TGC GTC GGA GGA ATG AAC AGA CGT CCA YNFMCNSSCVGGMNRR 961/321 991/331 ATT TTA ATC ATC GTT ACT CTG GAA ACC AGA GAT GGG CAA GTC CTG GGC CGA CGG TGC TTT I I V T L E T R D G Q V L G R R 1021/341 1051/351 gag gcc cgg atc tgt gcl tgc cca gga aga gac cgg aag gca gat gaa gac agc atc aga I C A C P G R D R K A D E D S 1081/361 1111/371 ang cag caa sta tog gac ago sca aas aac sec gat set acs aas cec cot tto cet cas V S D S A K N G D G T KRPFRQ 1141/381 1171/391 AAT ACA CAC GGA ATC CAG ATG ACT TCC ATC AAG AAA CGG AGA TCC CCA GAT GAT GAG CTG NTHGIQMTSIKKRRSPDD

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31/11
   ATG AAT TIT GAA ACT TOA CGG TGT GCC ACC CTA CAG TAC TGC CCC GAC CCT TAC ATC CAG
   M N F E T S R C A T L Q
                                       YCPDPYIQ
   61/21
                                  91/31
   CGT TTC ATA GAA ACC CCA GCT CAT TTC TCG TGG AAA GAA AGT TAT TAC AGA TCT GCC ATG
   RFIETPAHFSWKESYY
                                                   RSAM
                                  151/51
   TCG CAG AGC ACC CAG ACA AGC GAG TTC CTC AGC CCA GAG GTC TTC CAG CAT ATC TGG GAT
         S T Q T S E F
                                 SPEVFQHI
   181/61
                                 211/71
   TIT CTG GAA CAG CCT ATA TGC TCA GTA CAG CCC ATC GAG TTG AAC TIT GTG GAT GAA CCT
   F L E Q P I C S V Q P I E L N F V D E
                                 271/91
   TCC GAA AAT GOT GCa aCa aac aag att gag att agc atg gat tot atc coc atg caa gac
   S E N G A T N K I E I S M D C I R M Q
                                 331/111
   TCA GAC CTC AGT GAC CCC ATG TGG CCA CAG TAC ACG AAC CTG GGG CTC CTG AAC AGC ATG
   S D L S D P M W P Q
                                 Y T N L G L L N S M
   361/121
                                 391/131
  GAC CAG CAG ATT CAG AAC GGC TCC TCG TCC ACC AGC CCC TAC AAC ACA GAC CAC GCA CAG
  D Q Q I Q N G S S S T S P
                                         YNTDHAQ
421/141
                                451/151
  AAT AGC GTG ACG GCG CCC TCG CCC TAT GCA CAG CCC AGC TCC ACC TTT GAT GCC CTC TCT
N S V T A P S P Y A Q P S 5 T F D
E CCA TCC CCT GCC ATT CCC TCC AAC ACA GAT TAC CCG GGC CCA CAC AGC TTC GAT GTG TCC
P S P A I P S N T D Y P G P K S F
  541/181
TTC CAG CAG TCA AGC ACT GCC AAG TCA GCC ACC TGG ACG TAT TCC ACC GAA CTG AAG AAG
                                 571/191
F Q Q S S T A K S A T W T
[ 601/201
                                         YSTELKK
                                 631/211
  CTG TAC TGC CAG ATT GCG AAG ACA TGC CCC ATC CAG ATC AAG GTG ATG ACC CCA CCC CCA
  L Y C Q I A K T C P I Q I K V M
661/221
                                 691/231
CAG GGC GCT GTE ATC CGT GCC ATG CCT GTC TAC AAG AAA GCT GAG CAT GTC ACC GAG GTT
Q G A V I R A M P V Y K K A E H V T E V
                                751/251
GTG AAA CGA TGC CCT AAC CAT GAG CTG AGC CGT GAG TTC AAT GAG GGA CAG ATT GCC CCT
V K R C P N H E L S R E F N E G Q I
                                811/271
  CCC AGT CAT CTG ATT CGA GTA GAA GGG AAC AGC CAT GCC CAG TAT GTA GAA GAT CCT ATC
    S H L I R V E G N S H A Q Y V E D
  841/281
                                871/291
  ACG GGA AGG CAG AGC GTG CTG GTC CCT TAT GAG CCA CCA CAG GTT GGC ACT GAA TTC ACA
  T G R Q S V L V P Y E P P
  901/301
                                931/311
  ACA GTC CTG TAC AAT TTC ATG TGT AAC AGC AGC TGC GTC GGA GGA ATG AAC AGA CGT CCA
 T V L Y N F M C N S S C V G G M N R R P
                               991/331
 ATT TTA ATC ATC GTT ACT CTG GAA ACC AGA GAT GGG CAA GTC CTG GGC CGA CGG TGC TTT
       I I V T L E T R D G Q V L G R R C F
 GAG GCC CGG ATC TGT GCE TGC CCA GGA AGA GAC CGG AAG GCA GAT GAA GAC AGC ATC AGA
                                1051/351
    A R I C A C P G R D R K A D E D S I R
 1081/361
                                1111/371
 AAG CAG CAA GTA TCG GAC AGC GCA AAG AAC GGC GAT GGT ACG AAG CGC CCT TTC CGT CAG
          V S D S A K N G D G T K R P F R
 1141/381
                                1171/391
 AAT ACA CAC GGA ATC CAG ATG ACT TCC ATC AAG AAA CGG AGA TCC CCA GAT GAT GAG CTG
 N T H G I Q M T S I K K R R S P D D E L
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1201/401 1231/411 CTG TAC CTA CCA GTG AGA GGT GAG ACG TAC GAG ATG TTG CTG AAG ATG GAG TCA LYLPVRGRETYEMLLKI K E S 1261/421 1291/431 CTG GAG CTC ATG CAG TAC CTC CCT CAG CAC ACG ATC GAA ACG TAC AGG CAG CAG CAG LELMQYLPQHTIE T Y RQQQ 1351/451 CAG CAG CAC CAG CAC CTA CTT CAG AAA CAG ACC TCG ATG CAG TCT CAG TCT TCA TAT GGC Q Q H Q H L L Q K Q T S M Q S Q S S 1381/461 1411/471 1411/471 AAC AGT TCC CCA CCT CTG AAC AAA ATG AAC AGC ATG AAC AAG CTG CCT TCC GTG AGC CAG N S S P P L N K M N S M N K L P S V S Q 1441/481 1471/491 CTT ATC AAC CCA CAG CAG CGC AAT GCC CTC ACT CCC ACC ACC ATG CCT GAG GGC ATG GGA LINPQQRNAL TPT TMPEGM 1501/501 1531/511 GCC AAC ATT CCT ATG ATG GGC ACT CAC ATG CCA ATG GCT GGA GAC ATG AAT GGA CTC AGC ANIPMMGTHMPMAGDMNGL 1561/521 1591/531 CCT ACC CAA GCT CTC CCT CCT CCA CTC TCC ATG CCC TCC ACC TCC CAC TGC ACC CCA CCA PTQALPPPLSMPS TSHCTPP 1621/541 1651/551 CCG CCC TAC CCC ACA GAC TGC AGC ATT GTC AGG ATT tGG C&& GTC TGA PPYPTDCSIVRIWQV

23/42

31/11 ATG AAT TIT GAA ACT TCA CGG TGT GCC ACC CTA CAG TAC TGC CCC GAC CCT TAC ATC CAG MNFETSRCATLQYCPDP 91/31 CGT TTC ATA GAA ACC CCA GCT CAT TTC TCG TGG AAA GAA AGT TAT TAC AGA TCT GCC ATG RFIETPAHFSWKESYYR S 121/41 151/51 TCG CAG AGC ACC CAG ACA AGC GAG TTC CTC AGC CCA GAG GTC TTC CAG CAT ATC TGG GAT STQTSEFLSPEVFQH 181/61 211/71 TTT CTG GAA CAG CCT ATA TGC TCA GTA CAG CCC ATC GAG TTG AAC TTT GTG GAT GAA CCT F L E Q P I C S V Q P I E L N F V D E 241/81 271/91 TCC GAA AAT GGT GCa aCA aaC aaG ATT GAG ATT AGC ATG GAT TGT ATC CGC ATG Caa GAC SEN G A T N K I E I S M D C I R M 331/111 TCA GAC CTC AGT GAC CCC ATG TGG CCA CAG TAC ACG &AC CTG GGG CTC CTG &AC AGC ATG SDLSDPMWPQ YTN GLLN 361/121 391/131 GAC CAG CAG ATT CAG AAC GGC TCC TCG TCC ACC AGC CCC TAC BAC ACA GAC CAC GCA CAG D Q Q I Q N G S S S T S P YNTDHA 421/141 451/151 ANT AGO GTG ACG GCG CCC TCG CCC TAT GCA CAG CCC AGC TCC ACC TTL GAT GCC CTC TCT TAPSPYAQP55 TFDALS 511/171 CCA TGG GGT GCC ATT CCG TCC BAC ACA GAT TAC CCG GGC CCA CAC AGG TTC GAT GTG TCC S P A I P S N T D Y P G P H S F D 541/181 571/191 TTC CAG CAG TCA AGC AGT GCC AAG TCA GCC AGC TGG ACG TAT TCC ACC GAA CTG AAG AAG F Q Q S S T A K S A T W T Y S T E L K 601/201 631/211 CTG TAC TGC CAG ATT GCG AAG ACA TGC CCC ATC CAG ATC AAG GTG ATG ACC CCA CCC CCA LYCQIAKTCPIQIKVMTPP 661/221 691/231 CAG GGC GCT GTT ATC CGT GCC ATG CCT GTC TAC AAG AAA GCT GAG CAT GTC ACC GAG GTT QGAVIRAMPVYKKAEHVT 751/251 GTG AAA CGA TGC CCT AAC CAT GAG CTG AGC CGT GAG TTC AAT GAG GGA CAG ATT GCC CCT V K R C P N H E L S R E F N E G Q I A P 781/261 811/271 CCC AGT CAT CTG ATT CGA GTA GAA GGG AAC AGC CAT GCC CAG TAT GTA GAA GAT CCT ATC PSHLIRVEGNSHAQY VEDP 841/281 871/291 ACG GGA AGG CAG AGC GTG CTG GTC CCT TAT GAG CCA CCA CAG GTT GGC ACT GAA TTC ACA TGRQSVLVPYEPP 901/301 931/311 ACA GTC CTG TAC AAT TTC ATG TGT AAC AGC AGC TGC GTC GGA GGA ATG AAC AGA CGT CCA TVLYNFMCNSSCVGGMNR 961/321 991/331 ATT TTA ATC ATC GIT ACT CTG GAA ACC AGA GAT GGG CAA GTC CTG GGC CGA CGG TGC TTT I I V T L E T R D G Q LGRRC 1051/351 GAG GCC CGG ATC TGT GCT TGC CCA GGA AGA GAC CGG AAG GCA GAT GAA GAC AGC ATC AGA ARICACPGRDRK ADED 1081/361 1111/371 ANG CAG CAA GTA TCG GAC AGC GCA AAG AAC GGC GAT GCT TTC CGT CAG AAT ACA CAC GGA V S D S A K N G D A F R Q N T 1141/381 1171/391 ATC CAG ATG ACT TCC ATC AAG AAA CGG AGA TCC CCA GAT GAT GAG CTG CTG TAC CTA CCA I Q M T S I K K R R S P D D E L L Y L

1201/401 1231/411 OTG AGA GOT CGT GAG ACG TAC

V R G R E T Y

ATG TTG CTG AAG ATC AAA GAG TCA CTG

N L L K I K E S L ATG TTG CTG AAG ATC AAA GAG TCA CT CIC ATG 1261/421 1291/431 CAG TAC CTC CCT CAG CAC ACG ATC GAA ACG TAC AGG CAG CAG CAG CAG CAG CAG CAG Q Y L P Q H T I E T Y R Q Q Q Q Q R Q 1321/441 1351/451 CAC CTA CTT CAG AAA CAT CTC CTT TCA GCC TGC TTC AGG AAT GAG CTT GTG GAG CCC CGG H L L Q K H L L S A C F R N E L V E P R 1381/461 1411/471 GGA GAA GCT CCG ACA CAG TCT GAC GTC TTC TTT AGA CAT TCC AAC CCC CCA AAC CAC TCC G E A P T Q S D V F F R H S N P P N H S 1441/481 GTG TAC CCA TAG V Y P +

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31/11
   ATG TTG TAC CTG GAA AAC AAT GCC CAG ACT CAA TTT AGT GAG CCA CAG TAC ACG AAC CTG
   M L
         YLENNAQTQFSEPQYTNL
   61/21
                                 91/31
   GGG CTC CTG AAC AGC ATG GAC CAG CAG ATT CAG AAC GGC TCC TCG TCC ACC AGC CCC TAC
   G L L N S M D Q Q I
                                 Q N G
                                         S S
                                               S
   121/41
                                 151/51
   ARC ACA GAC CAC GCA CAG AAT AGC GTG ACG GCG CCC TCG CCC TAT GCA CAG CCC AGC TCC
   NTDHAQNSVTAPSPYAQPS
   181/61
                                 211/71
   ACC TIT GAT GCC CTC TCT CCA TCC CCT GCC ATT CCC TCC AAC ACA GAT TAC CCG GGC CCA
   T F D A L S P S P A I P S N T D Y P
                                 271/91
   CAC AGC TTC GAT GTG TCC TTC CAG CAG TCA AGC ACT GCC AAG TCA GCC ACC TGG ACG TAT
        F D V. S F Q Q S S T A K S A T W
   301/101
                                 331/111
   TCC ACC GAA CTG AAG AAG CTG TAC TGC CAG ATT GCG AAG ACA TGC CCC ATC CAG ATC AAG
   S T E L K K L Y C Q I A K T C P I Q
   361/121
                                 391/131
   GTG ATG ACC CCA CCC CCA CAG GGC GCT GTL ATC CGT GCC ATG CCT GTC TAC AAG AAA GCT
   V M T P P P Q G A V
                                I R A
                                         MPVYKKA
   421/141
                                 451/151
   GAG CAT GTC ACC GAG GTT GTG AAA CGA TGC CCT AAC CAT GAG CTG AGC CGT GAG TTC AAT
   E H V T E V V K R C P N H E L S R E
   481/161
                                 511/171
   GAG GGA CAG ATT GCC CCT CCC AGT CAT CTG ATT CGA GTA GAA GGG AAC AGC CAT GCC CAG
   EGQIAPPSHL
                                 IRVEGNSHAQ
   541/181
                                 571/191
   TAT GTA GAA GAT CCT ATC ACG GGA AGG CAG AGC GTG CTG GTC CCT TAT GAG CCA CCA CAG
   YVEDPITGRQSVL
                                         VPYEP
   601/201
                                631/211
  GTT GGC ACT GAA TTC ACA ACA GTC CTG TAC AAT TTC ATG TGT AAC AGC AGC TGC GTC GGA
     G T E F T T V L Y N F M C N S
  661/221
                                 691/231
  GGA ATG AAC AGA CGT CCA ATT TTA ATC ATC GTT ACT CTG GAA ACC AGA GAT GGG CAA GTC
  G M N R R P I L I I
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                                V T L E T R D G Q
  721/241
                                 751/251
  CTG GGC CGA CGG TGC TTT GAG GCC CGG ATC TGT GCL TGC CCA GGA AGA GAC CGG AAG GCA
  LGRRCFEARICAC
                                         PGRDRKA
  781/261
                                 811/271
  gat gaa gac agc atc aga aag cag caa gta tog gac agc gca aag aac ggc gat ggt acg
          S I R K Q Q V S D S A K N G D G T
                                871/291
  AAG CGC CCT TTC CGT CAG AAT ACA CAC GGA ATC CAG ATG ACT TCC ATC AAG AAA CGG AGA
        PFRQNTHGIQMTSIKKR
  901/301
                                931/311
  TCC CCA GAT GAT GAG CTG CTG TAC CTA CCA GTG AGA GGT CGT GAG ACG TAC GAG ATG TTG
  S P D D E L L Y L P V R G R E T Y E M L
  961/321
                                991/331
  CTG AAG ATC AAA GAG TCA CTG GAG CTC ATG CAG TAC CTC CCT CAG CAC ACG ATC GAA ACG
     K I K E S L E L M Q Y L P Q H T I
  1021/341
                                1051/351
  TAC AGG CAG CAG CAG CAG CAG CAG CAG CTA CTT CAG AAA CAG ACC TCG ATG CAG
  YRQQQQQRHQHLLQXQT
  1081/361
                                1111/371
  TCT CAG TCT TCA TAT GGC AAC AGT TCC CCA CCT CTG AAC AAA ATG AAC AGC ATG AAC AAG
          SYGNSSPPLN
                                        K M N S M N
  1141/381
                                1171/391
  CTG CCT TCC GTG AGC CAG CTT ATC AAC CCA CAG CAG CGC AAT GCC CTC ACT CCC ACC ACC
  L P S V S Q L I N P Q Q R N A L T P T T
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31/11

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ATG TTG TAC CTG GAA AAC AAT GCC CAG ACT CAA TTT AGT GAG CCA CAG TAC ACG AAC CTG
    M L Y L E N N A Q T Q F S E P Q Y T N
                                  91/31
    GGG CTC CTG AAC AGC ATG GAC CAG ATT CAG AAC GGC TCC TCG TCC ACC AGC CCC TAC
    G L L N S M D Q Q I
                                 QNGSSST
    121/41
                                  151/51
    ARC ACA GAC CAC GCA CAG AAT AGC GTG ACG GCG CCC TGG CCC TAT GCA CAG CCC AGC TCC
    NTDHAQNSVTAP
                                          PYAQPSS
    181/61
                                  211/71
    ACC TTT GAT GCC CTC TCT CCA TCC CCT GCC ATT CCC TCC AAC ACA GAT TAC CCG GGC CCA
    TFDALSPSPA
                                 I P S N T D Y P
    241/81
                                 271/91
    CAC AGC TTC GAT GTG TCC TTC CAG CAG TCA AGC ACT GCC AAG TCA GCC ACC TGG ACG TAT
    H S F D V S F Q Q S S T A
                                          KSATWT
                                 331/111
    TCC ACC GAA CTG AAG AAG CTG TAC TGC CAG ATT GCG AAG ACA TGC CCC ATC CAG ATC AAG
    S T E L K K L Y C Q
                                IAKTCP
    361/121
                                 391/131
    GTG ATG ACC CCA CCC CCA CAG GGC GCT GTL ATC CGT GCC ATG CCT GTC TAC AAG AAA GCT
         T P P P Q G A V
                                 I R A M P V Y K K A
    421/141
                                 451/151
   GAG CAT GTC ACC GAG GTT GTG AAA CGA TGC CCT AAC CAT GAG CTG AGC CGT GAG TTC AAT
   E H V T E V V K R C P N H E L S R E F
                                 511/171
   GAG GGA CAG ATT GCC CCT CCC AGT CAT CTG ATT CGA GTA GAA GGG AAC AGC CAT GCC CAG
   E G Q I A P P S H L I R V E G N S
                                 571/191
   TAT GTA GAA GAT CCT ATC ACG GGA AGG CAG AGC GTG CTG GTC CCT TAT GAG CCA CCA CAG
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   YVEDPITGRQSVLVPYEPP
ŀĀ
                                 631/211
   GTT GGC ACT GAA TTC ACA ACA GTC CTG TAC BAT TTC ATG TGT AAC AGC AGC TGC GTC GGA
   V G T E F T T V L Y N F M C N S S C V G
                                691/231
   GGA ATG AAC AGA CGT CCA ATT TTA ATC ATC GTT ACT CTG GAA ACC AGA GAT GGG CAA GTC
     MNRRPILIIVTLETRDGQ
   721/241
   CTG GGC CGA CGG TGC TTT GAG GCC CGG ATC TGT GCL TGC CCA GGA AGA GAC CGG AAG GCA
                                751/251
   LGRRCFEARICACPGRDRKA
                                811/271
   GAT GAA GAC AGC ATC AGA AAG CAG CAA GTA TCG GAC AGC GCA AAG AAC GGC GAT GGT ACG
  D E D S I R K Q Q V S D S A K N G D G
                                871/291
   ARG CGC CCT TTC CGT CAG ART ACA CAC GGA ATC CAG ATG ACT TCC ATC AAG AAA CGG AGA
  K R P F R Q N T H G I Q M T S I K K R R
   TCC CCA GAT GAT GAG CTG CTG TAC CTA CCA GTG AGA GGT CGT GAG ACG TAC GAG ATG TTG
                                931/311
    b D D E T T A T b
                               VRGRET
                                                 YEML
                                991/331
  CTG AAG ATC AAA GAG TCA CTG GAG CTC ATG CAG TAC CTC CCT CAG CAC ACG ATC GAA ACG
     KIKESLELMQYLPQHTI
                                1051/351
  TAC AGG CAG CAG CAG CAG CAG CAG CAG CTA CTT CAG AAA CAG ACC TCG ATG CAG
  YRQQQQQHQHLLQXQTSMQ
  TCT CAG TCT TCA TAT GGC AAC AGT TCC CCA CCT CTG AAC AAA ATG AAC AGC ATG AAC AAG
                                1111/371
         S Y G N S S P P L N K M N S M N
  1141/381
  CTG CCT TCC GTG AGC CAG CTT ATC AAC CCA CAG CAG CGC AAT GCC CTC ACT CCC ACC ACC
          V S Q L I N P Q Q R N A L T P T
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31/11 ATG TTG TAC CTG GAA AAC AAT GCC CAG ACT CAA TTT AGT GAG CCA CAG TAC ACG AAC CTG MLYLENNAQTQFSEPQYT 61/21 91/31 GGG CTC CTG AAC AGC ATG GAC CAG CAG ATT CAG AAC GGC TCC TCG TCC ACC AGC CCC TAC G L L N S M D Q Q I Q N G S S s T 151/51 121/41 AAC ACA GAC CAC GCA CAG AAT AGC GTG ACG GCG CCC TCG CCC TAT GCA CAG CCC AGC TCC N T D H A Q N S V T APSPYAQ 181/61 211/71 ACC TTT GAT GCC CTC TCT CCA TCC CCT GCC ATT CCC TCC AAC ACA GAT TAC CCG GGC CCA TFD'ALSFSPA I P S N T D Y P G 271/91 CAC AGC TTC GAT GTG TCC TTC CAG CAG TCA AGC ACT GCC AAG TCA GCC ACC TGg ACG TAT H S F D V S F Q Q S S T A WTAZX 301/101 331/111 TCC ACC GAA CTG AAG AAG CTG TAC TGC CAG ATT GCG AAG ACA TGC CCC ATC CAG ATC AAG S T E L K K L Y C Q I A K T C P I Q 361/121 391/131 GTG ATG ACC CCA CCC CCA CAG GGC GCT GTT ATC CGT GCC ATG CCT GTC TAC AAG AAA GCT V M T P P P Q G A V I R A MPVYKK 421/141 451/151 GAG CAT GTC ACC GAG GTT GTG AAA CGA tGC CCT AAC CAT GAG CTG AGC CGT GAG TTC AAT EHVTEVVKRCPNHELSREFN 511/171 GAG GGA CAG ATT GCC CCT CCC AGT CAT CTG ATT CGA GTA GAA GGG AAC AGC CAT GCC CAG EGQIAPPSHL I R V EGNSHA 541/181 571/191 THE GIA GAR GAT COT ATC ACG GGA AGG CAG AGC GTG CTG GTC CCT LAT GAG CCA CCA CAG V E D P ITGRQSVLV 601/201 631/211 GTT GGC ACT GAA TTC ACA ACA GTC CTG TAC AAT TTC ATG TGT BAC AGC AGC TGC GTC GGA V G T E F T T V L Y N F M C N S S C V G 661/221 691/231 GGA ATG AAC AGA CGT CCA ATT TTA ATC ATC GTT ACT CTG GAA ACC AGA GAT GGG CAR GTC G M N R R F I L I I V T L E T R D G Q V 721/241 751/251 CTG GGC CGA CGG TGC TTT GAG GCC CGG ATC TGT GCT TGC CCA GGA AGA GAC CGG AAG GCA LGRRCFEARICACF GRDR 781/261 811/271 GAT GAA GAC AGC ATC AGA AAG CAG CAA GTA TCG GAC AGC GCA AAG AAC GGC GAT GCT TTC D E D S I R K Q Q V SDSAKNGDA 841/281 871/291 COT CAG AAT ACA CAC GGA ATC CAG ATG ACT TCC ATC AAG AAA CGG AGA TCC CCA GAT GAT RQNTHGIQMT SIKKRRSPDD 901/301 931/311 gag ctg ctg tac cta cca gtg aga ggt cgt gag acg tac gag atg ttg ctg aag atc aaa ELLYLPVRGRETYEMLLK 961/321 991/331 gag tea etg gag etc atg eag tac etc eet eag eac acg ate gaa acg tac agg eag eag L E L M Q Y L P QHT 1021/341 1051/351 CAG CAG CAG CAG CAG CAG CTA CTT CAG AAA CAT CTC CTT TCA GCC TGC TTC AGG AAT QQQHQHLLQKHLLSACFRM 1081/361 1111/371 GAG CTT GTG GAG CCC CGG GGA GAA GCT CCG ACA CAG TCT GAC GTC TTC TTT AGA CAT TCC V E P R G E A P T Q S D V F F R H S 1141/381 AAC CCC CCA AAC CAC TCC GTG TAC CCA TAG N P P N H S V Y P

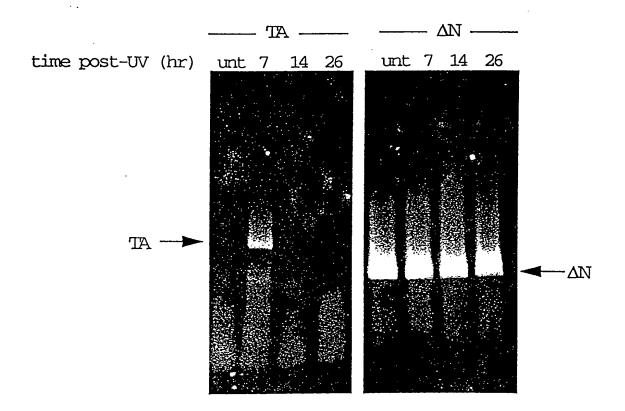


Figure 21. UV Induction of TA-p63 transcript. Primary human foreskin keratinocytes were exposed to UV irradiation ($300J/m^2$), then harvested at times indicated. Total RNA was prepared, and RT-PCR on equal amounts of RNA was performed using primers specific for transactivating (TA) or truncated (Δ N) N-terminal domains of p63 (Yang et al., 1998). TA-p63 is not detectable in untreated (unt) keratinocytes, but expression is induced at 7hrs after UV treatment. Δ N-p63 appears to remain unchanged from untreated cells at all time points taken.

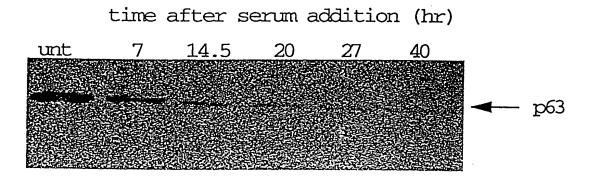


Figure 22. p63 protein expression decreases upon keratinocyte differentiation. Primary human foreskin keratinocytes were treated with 10% fetal bovine serum to induce differentiation, and harvested at times indicated. Cells were lysed in RIPA buffer, boiled in Lammeli buffer, and proteins fractionated by SDS gel electrophoresis. Immunoblotting was performed using an anti-p63 mouse monoclonal antibody, 4A4, which recognizes all p63 isoforms identified to date. p63 protein levels in keratinocytes are seen to decrease progressively upon differentiation, as compared to untreated (unt) samples.

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Figure 23. p63 RNA expression in human cancer cell lines. RT-PCR was performed on total RNA from several human tumor cell lines* using primers specific for transactivating (TA) or truncated (Δ N) N-terminal domains of p63 (see Figure 1), as indicated. Amongst these cell lines, only the ME180 control cell line showed a TA-p63 transcript. Instead, a majority showed expression of Δ N-p63, which has been demonstrated to act as a dominant negative protein towards both the tumor suppressor p53, as well as transactivating isotypes of p63.

^{*}note: all are cervical carcinoma cell lines except U2OS, which is a human osteocarcinoma line.

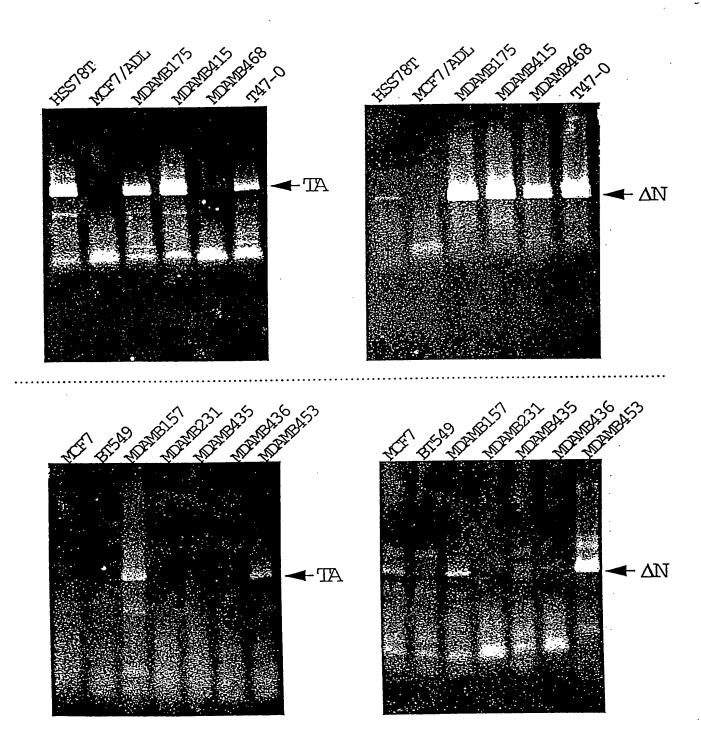
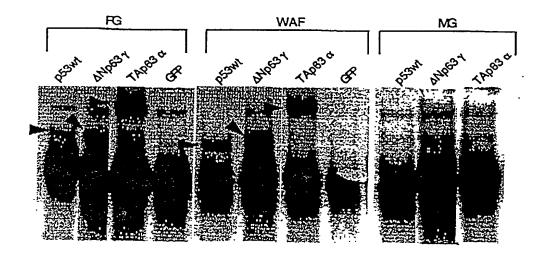


Figure 24. p63 RNA expression in human breast cancer cell lines. RT-PCR was performed on total RNA from breast cancer cell lines using primers specific for transactivating (TA) or truncated (ΔN) N-terminal domains of p63 (see Figure 1), as indicated. p63 was detected in a majority of cell lines tested, with some specimens containing both transcript isotypes.

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Electrophoretic mobility shift assays (EMSA) were performed using three separate, ³²P radiolabeled oligonucelotides: a minimal p53 binding sequence site (PG), a p53 binding site in the p21 promoter (WAF), and a mutant p53 binding site (MG, *Kern et al., 1992) with lysates of 293 human kidney cells transfected with p53, ΔNp63γ, TAp63α, and green fluorescent protein (GFP). p53, ΔNp63γ, and TAp63α lysates all yielded significant mobility shifts of both PG and WAF oligonucleotides (highlighted by arrows), while the GFP protein, included as negative control, failed to display a similar shift. None of the lysates showed a shift of the control, non-p53 binding oligonucleotide, MG, thus demonstrating the specificity of p53, ΔNp63γ, and TAp63α interactions with the p53-binding sites.

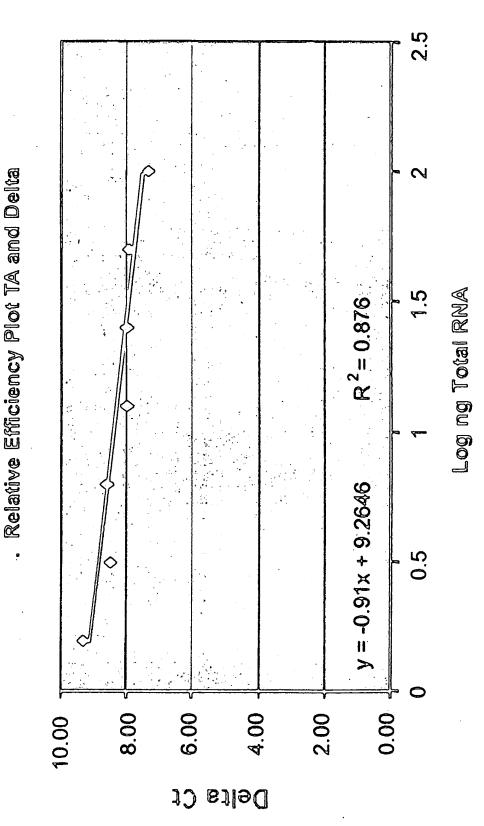


FIGURE 26

A. TAp63 amplification plot for PrEC cells

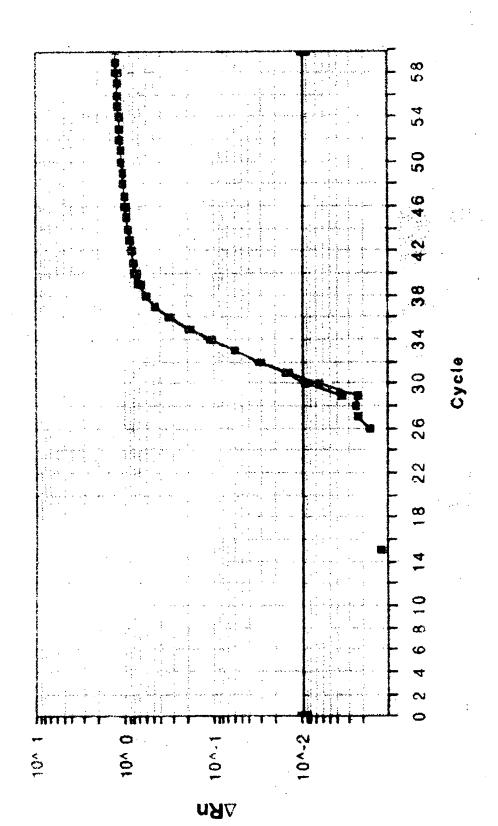
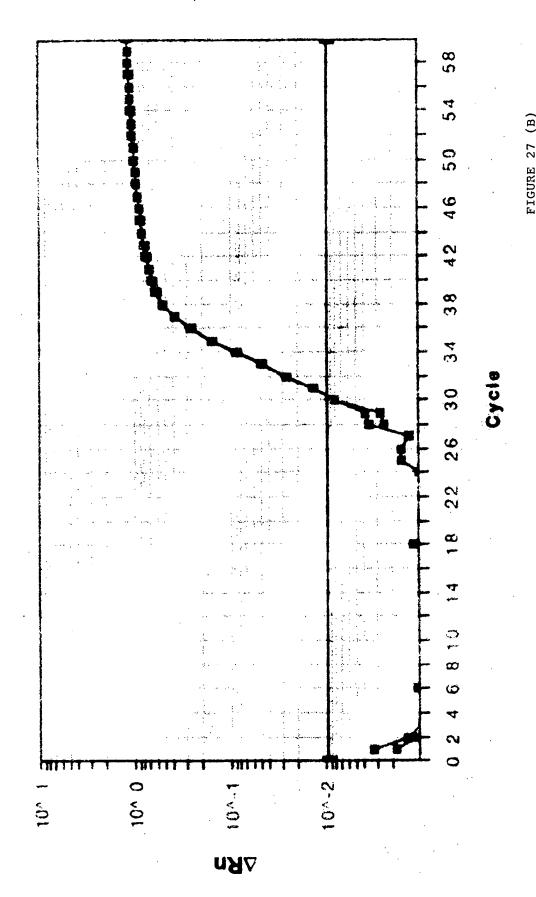


FIGURE 27(A)

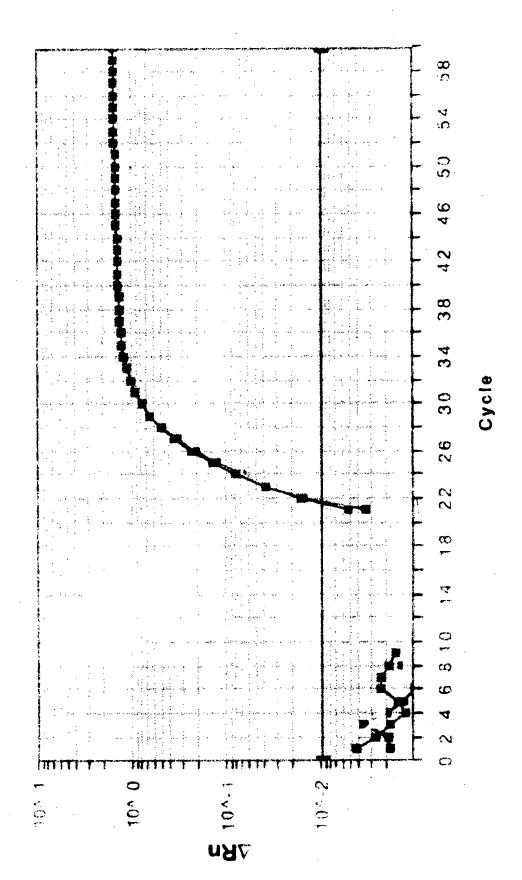
B. TAp63 amplification plot for PC3 cells



C. ANp63 amplification plot for PrEC cells

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D. ANp63 amplification plot for PC3 cells

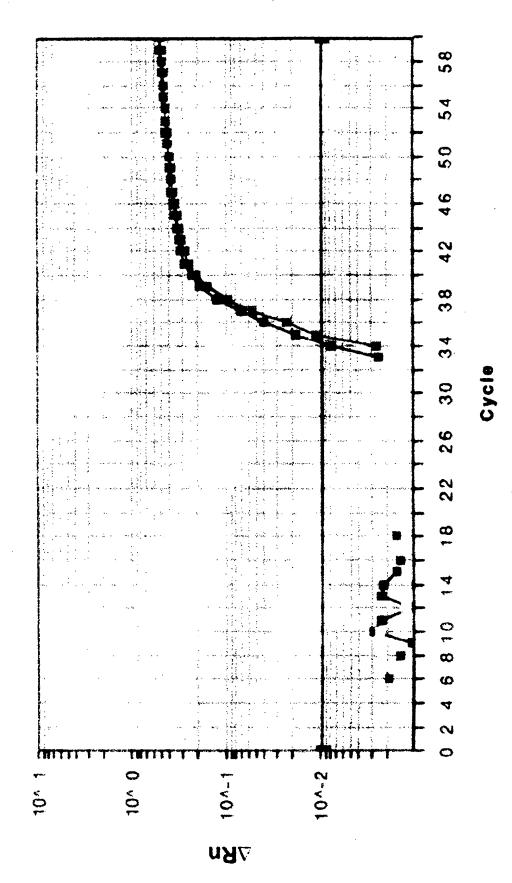
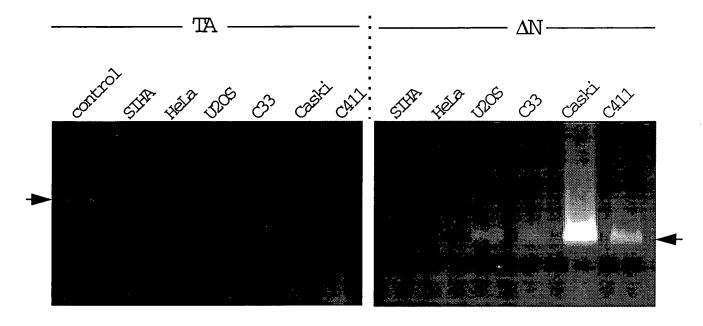


FIGURE 27(D)



p63 RNA expression in human cancer cell lines. RT-PCR was performed on total RNA from several human tumor cell lines* using primers specific for transactivating (TA) or truncated (Δ N) N-terminal domains of p63 as indicated. Amongst these cell lines, only theME180 control cell line showed a TA-p63 transcript. Instead, a majority showed expression of Δ N-p63, which has been demonstrated to act as a dominant negative protein towards both the tumor suppressor p53, as well as transactivating isotypes of p63.

^{*}note: all are cervical carcinoma cell lines except U2OS, which is a human osteocarcinoma line.

4A4 anti-p63

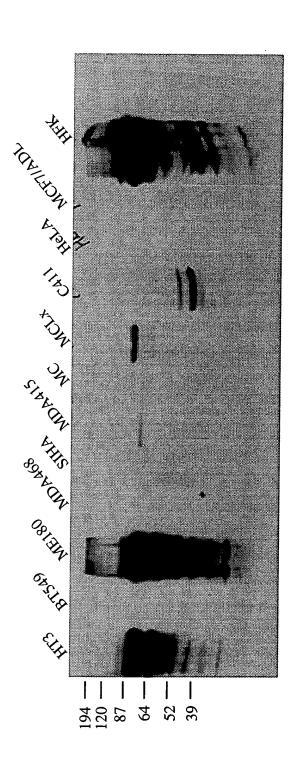


FIGURE 29